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RECEIVED 18 November 2024 ACCEPTED 21 March 2025 PUBLISHED 24 April 2025

CITATION

Xu Z, Tao Z and Guo Y (2025) The role of tea in managing cardiovascular risk factors: potential benefits, mechanisms, and interventional strategies. *Front. Nutr.* 12:1530012. doi: 10.3389/fnut.2025.1530012

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The role of tea in managing cardiovascular risk factors: potential benefits, mechanisms, and interventional strategies

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Traditional major cardiovascular disease (CVD) risk factors include dyslipidemia, hypertension, smoking, diabetes, and obesity. Tea is rich in various bioactive substances such as tea polyphenols, theaflavins, and tea polysaccharides. Due to the regulatory effects on multiple pathways and its anti-inflammatory and antioxidant properties, these active substances have shown significant efficacy in regulating dyslipidemia, hypertension, diabetes, obesity, and cardiac autonomic function. Additionally, tea possesses anti-inflammatory and antithrombotic properties, making it a promising dietary supplement for nutritional interventions in the primary and secondary prevention of CVDs. However, the complex composition of tea, although shown to have certain effects *in vivo*, does not fully elucidate the specific mechanisms of action. Moreover, the varying application methods across different studies lead to differences in intervention effects and dose–response relationships, sometimes resulting in contradictory findings. This article reviews the potential benefits, mechanisms of action, and application methods of tea for cardiovascular risk factors, elucidating its potential as a nutritional intervention.

KEYWORDS

tea, cardiovascular risk factors, non-pharmacological treatment, nutritional intervention, interventional strategies

1 Introduction

Cardiovascular diseases (CVDs) are significantly influenced by several risk factors, including dyslipidemia, hypertension, smoking, diabetes mellitus, and obesity (1). These factors are widely recognized not only in adult populations but are increasingly prevalent among children and adolescents. This shift has resulted in a higher incidence of early atherosclerosis and cardiovascular events (2, 3).

Obesity, in particular, exhibits a close relationship with other cardiovascular risk factors such as hypertension, diabetes, and hyperlipidemia, thereby forming a complex and interconnected network of risk factors (4). Additionally, smoking is a critical behavioral risk factor that further elevates the risk of CVDs, especially in individuals with other metabolic abnormalities (5).

Therefore, the early identification and management of these risk factors are essential to mitigate the incidence and mortality associated with cardiovascular diseases (6). Recent research has also identified additional potential independent risk factors, including gender differences, aging, and psychosocial factors, which warrant further exploration and consideration (7, 8).

The management of these risk factors includes both pharmacological and non-pharmacological approaches. Pharmacological treatments encompass lipid regulation,

blood pressure control, diabetes treatment, as well as symptomatic treatment for other independent risk factors. For patients already diagnosed with atherosclerotic cardiovascular diseases, antiinflammatory and antithrombotic interventions are also considered crucial for controlling underlying risk factors.

Non-pharmacological treatments include lifestyle modifications, nutritional interventions, physical activity, and psychological interventions. The importance of non-pharmacological treatments has been increasing in light of changes in social environment, shifts in health awareness, and the continuous development of adjunctive therapies. Numerous studies have confirmed that effective management of these risk factors can significantly improve disease symptoms and reduce the recurrence and rehospitalization rates of CVD, thereby extending patient life expectancy and improving quality of life.

Nutritional interventions cover four main areas: dietary structure, obesity and weight management, fat and sugar intake, and the use of dietary supplements. The goal is to prevent and actively improve cardiovascular diseases through nutritional diets. Tea, as a soft drink, is rich in phenols, polysaccharides, and flavonoids, and has high nutritional and health value. These components have been shown to have positive effects on reducing the risk and progression of CVD, making tea a suitable dietary supplement for cardiovascular disease needs (9). Tea can be classified into six categories based on the degree of fermentation: green tea, white tea, yellow tea, oolong tea, black tea, and dark tea. A large body of clinical and mechanistic research supports that the chemical components in tea have positive effects on lipids, blood pressure, diabetes, obesity, inflammation, and thrombosis.

2 Chemical composition of tea

Tea contains various chemical constituents including tea polyphenols, tea pigments, tea polysaccharides, tea saponins, alkaloids, aromatic substances, vitamins, amino acids, and a small amount of inorganic substances. Tea polyphenols and tea polysaccharides are considered the primary sources of the health benefits of tea, and their content varies depending on the type and degree of fermentation.

Catechins are the most significant chemical constituents of tea polyphenols, accounting for about 60–80% of the total tea polyphenols. They can be divided into four categories: epigallocatechin gallate (EGCG), epigallocatechin (EGC), epicatechin gallate (ECG), and epicatechin (EC) (10). During tea fermentation, some catechins are oxidized to theaflavins, resulting in a decrease in the total catechin content (from approximately 172.8 mg/g to 48.2 mg/g), and an increase in theaflavins (from approximately 17.9 mg/g to 43.4 mg/g) (11). The antioxidant activity of theaflavins is lower than that of catechins and includes four isomers: theaflavin (TF1), theaflavin-3-gallate (TF2A), theaflavin-3'-gallate (TF2B), and theaflavin-3,3'-digallate (TF3) (12).

Tea polysaccharides are non-starch bound acidic polysaccharides composed of neutral sugars, uronic acids, and proteins. The monosaccharide components of tea polysaccharides primarily include rhamnose, arabinose, xylose, mannose, glucose, galactose, and fucose (13). The variation in the monosaccharide composition and content, as well as differences in polysaccharide structures across different types of tea, suggest that tea polysaccharides are unique components of tea (Figure 1).

3 Benefits and mechanisms of tea on cardiovascular risk factors

3.1 Lipids

Dyslipidemia is considered a critical factor in atherosclerosis and related cardiovascular events. The association between low-density lipoprotein and other apolipoprotein B-containing lipoproteins and



cardiovascular event risk has also been confirmed through Mendelian randomization studies in humans (14). Moreover, studies have shown that tea consumption not only significantly improves hyperlipidemia but also reduces the risk of coronary atherosclerotic heart disease (15, 16).

Early studies attributed the lipid metabolism-enhancing effects of tea to the role of caffeine in promoting fat breakdown (17). However, recent findings demonstrate a significant positive correlation between the active components of tea and its lipid-lowering and antioxidant properties. These effects may be associated with increased excretion of bile acids and cholesterol, as well as enhanced activities of catalase and glutathione peroxidase (18). Cross-sectional studies indicate that women aged 20 to 48 derive higher average antioxidant capacity and anti-lipid oxidative damage benefits from dietary intake of total flavonoids and theaflavins in tea compared to vitamins and anthocyanins (19). Raederstorff et al. observed a reduction in intestinal cholesterol absorption rates in rats from 73.7 to 62.7% when exposed to an intervention of 0.5 g/kg epigallocatechin gallate (EGCG) compared to 0.1 g/kg. Correspondingly, in vitro experiments demonstrated that EGCG dose-dependently decreased cholesterol solubility in micelles, suggesting that it may improve lipid metabolism by inhibiting the solubilization of cholesterol in the digestive tract and reducing its absorption (20). Furthermore, thearubigins have been shown to elevate conjugated bile acid levels, suppress the intestinal FXR-FGF15 signaling pathway, reduce cholesterol and fat synthesis, and activate alternative bile acid synthesis pathways to promote fat breakdown (21). Kashif et al. examined the effects of catechins, theaflavins, and freeze-dried ginger extracts in hyperglycemic, obese, and liver-impaired rats. Their findings revealed that catechins significantly improved body weight, cholesterol (-11.03%), and low-density lipoprotein (LDL) (-14.25%), with enhanced results when combined with theaflavins (22). Fu et al. (131) further identified that theaflavin (TFDG) lowered fasting blood glucose and lipid concentrations by upregulating the Nrf2 signaling pathway and circ-ITCH expression. Their research validated the antioxidant properties of TFDG and its beneficial effects on liver and kidney function, as well as cellular structural integrity (23).

While research has shown that tea and its components can improve blood lipid levels, combining tea with other interventions may result in enhanced effects (24, 25). However, the relationship between tea-based intervention strategies and the regulation of blood lipids remains an area of significant interest. Elke et al. (132) conducted a study involving 102 participants with mild-to-moderate hypercholesterolemia (TC: 5.70 ± 0.74 and/or LDL-C: 3.97 ± 0.61 mmol/L) using tea components and cellulose as interventions. Although total cholesterol (TC) and low-density lipoprotein cholesterol (LDL-C) levels decreased over the intervention period, no statistically significant difference was observed compared to the placebo group. It was concluded that the lipid-lowering effects of tea could not be confirmed at a daily dosage of 75 mg theaflavins and 150 mg catechins, whether applied individually or in combination (26). Patricia et al. (133) found in their study on type 2 diabetes patients undergoing standard therapy and statin treatment that a 12-week intervention with 400 mg green tea extract (90% total polyphenols, 80% total catechins, 45% EGCG, and 1.0% caffeine) did not significantly alter blood lipid or glucose levels compared to placebo. However, the green tea extract positively impacted arterial health by reducing the central arterial augmentation index $(-3.05 \pm 10.8\%$ vs. 6.7 \pm 0.1%), suggesting an improvement in arterial stiffness (27). The dosages of tea components applied in the above studies showed no significant intervention effects; however, alternative dosages or the use of brewed tea were not further investigated. In contrast, Bianca et al. (134) evaluated a six-week intervention involving combined supplementation of fish oil (1.7 g EPA + DHA/day), plant sterol-enriched chocolate (2.2 g/day), and green tea (~170.8 mg/day) in 53 statin-intolerant type 2 diabetes patients. Responders (n = 10) continued with a 12-week statin dose-reduction combined with supplementation. No significant differences were observed in lipid levels or inflammatory response improvements compared to standard therapy (28). Shun et al. (135), in a double-blind, placebo-controlled study, demonstrated that a four-week intervention with 165 mg mono-glucosyl hesperidin and 387 mg green tea catechins significantly reduced triglyceride (TG) levels (29).

In summary, tea's lipid-improving effects may be more pronounced when combined with pharmaceutical or supplementary interventions, rather than relying solely on tea or its extracts. Independent application of tea extracts may not achieve optimal outcomes for blood lipid regulation (Figure 2).

3.2 Blood pressure

Hypertension is a common cardiovascular disease characterized by elevated blood pressure. It causes target organ damage through hemodynamic changes and, in conjunction with other risk factors, increases the risk of cardiovascular diseases and events (30). Retrospective studies by Jonathan et al. (136) observed that long-term regular tea consumption led to an average reduction of 2-3 mmHg in systolic and diastolic blood pressure in elderly women, with particularly notable effects at a daily intake of 250 mL. However, the study did not specify the actual concentration or type of tea consumed (31). Similarly, a study conducted in Taiwan involving 1,507 tea-drinking participants examined the impact of tea consumption on the risk of developing hypertension. The results revealed that habitual moderate tea consumption (120 mL/day for more than one year) significantly reduced the incidence of hypertension (32). Furthermore, a meta-analysis of randomized controlled trials demonstrated that tea components reduced oxidative stress and inflammatory responses while improving vasodilation associated with blood pressure changes (33). These findings provide supporting evidence for the long-term use of tea as a potential strategy for the prevention and management of hypertension.

Furthermore, clinical research shows that acute tea consumption significantly increases systolic and diastolic blood pressure after 120 min, although subjects showed the lowest digital volume pulse stiffness index (34). Jonathan et al. (136) also confirmed that acute tea consumption raised systolic blood pressure within three hours (35). The acute rise in blood pressure from tea consumption may be due to caffeine, but long-term tea consumption has a positive effect on blood pressure regulation, potentially related to polyphenolic compounds (Figure 3).

Hyperhomocysteinemia-associated hypertension is gaining increasing attention in clinical diagnosis and treatment, as it is also a risk factor for atherosclerotic cardiovascular disease (36). Long-term tea consumption has been shown to improve endothelial function, reduce hemodynamic-mediated vascular dilation, and thereby improve blood pressure levels. These effects may be associated with the inhibition of angiotensin by tea pigments and theaflavins (37–39). Lili et al. (137)





FIGURE 3

Chemical content of tea (93) (Sorted by white tea, green tea, yellow tea, oolong tea, black tea, and black tea categories. **White tea** includes Gongmei white tea, White Peony tea; **green tea** includes Dianqing tea, Dongting Biluochun tea, Duyun Maojian tea, Enshi Yulu tea, Lu'an Guapian tea, Lushan Yunwu tea, Taiping Houkui tea, Xihu Longjing tea, Yongxi Huoqing tea; **yellow tea** includes Huoshan large yellow tea, Junshan Yinzhen tea, Mengding Huangya tea, Weishan Maojian tea, Yuan'an Luyuan tea; **oolong tea** includes Fenghuang Shuixian tea, Luohan Chenxiang tea, Tieguanyin tea, Wuyi Rock tea; **black tea** includes Dianhong Congou black tea, Keemun black tea, Lapsang Souchong black tea, Yichang Congou black tea; and **dark tea** includes Fuzhuan Brick tea, Liupao tea, Pu-erh tea, Qingzhuan Brick tea, Tibetan tea).

demonstrated that epigallocatechin gallate (EGCG), due to its potential antioxidant and anti-inflammatory activities, significantly improved cerebrovascular damage in rats induced by high Hcy levels. It also increased glutathione levels, enhancing antioxidant capacity (40). Tea polyphenols and bioflavonoids were found to inhibit DNMT-mediated DNA methylation, which partially restored HUVEC cell damage induced by high Hcy levels and reduced PAI-1 activity (41, 42). Notably, the risk of high Hcy levels may be positively correlated with the intake of black or green tea due to individual differences in the methylation of polyphenolic compounds. In contrast, oolong tea consumption was not associated with an increased risk of high Hcy levels. Moreover, theaflavins in oolong tea exhibited protective effects against Hcy-induced endothelial damage (43–45).

Hypertension and cellular damage are closely linked, and their treatment are crucial for improving cardiovascular diseases (46). Catechins, major antioxidant components of tea polyphenols, improve oxidative stress in cardiomyocytes by reducing reactive oxygen species production in cells and mitochondria and decreasing antioxidant factor consumption. This process may depend on the inhibition of inflammation-related pathways, leading to a reduction in interleukin-8 (IL-8) production (47). Aravind et al. (138) demonstrated that EGCG not only inhibited the NF-kB pathway, reducing the transcription of downstream inflammatory factor genes, but also decreased type II coronary endothelial cell extravasation and monocyte adhesion. This blocked cell activation and further validated the positive role of EGCG in mitigating inflammatory responses and maintaining vascular homeostasis (48).

The regulatory effects of tea on blood pressure vary across studies, which may be closely linked to differences in tea types and intervention dosages. Marjan et al. (139) conducted a meta-analysis of five RCT studies involving 408 participants, finding that habitual tea consumption reduced SBP and DBP by approximately -3.53 mmHg and - 0.99 mmHg, respectively. The blood pressure reduction was more pronounced with tea consumption durations of \geq 3 months, with green tea showing greater efficacy in lowering blood pressure compared to black tea (49). Additionally, Biesinger et al. (140) used a combination of phytochemicals (grape seed and skin extract 330 mg, green tea 100 mg, resveratrol 60 mg, and a mixture of quercetin, ginkgo, and bilberry 60 mg) for placebo-controlled crossover intervention in metabolic syndrome subjects with elevated blood pressure, finding that phytochemical supplements reduced mean arterial pressure and diastolic pressure by 4.4 mmHg. Although participants' blood pressure improved in this study, the complexity of the chemical combination used means the improvement cannot be solely attributed to green tea (50). A clinical randomized crossover trial found that taking three capsules of 500 mg green tea extract (260 mg polyphenols) daily for four weeks significantly reduced 24-h systolic blood pressure (-3.61 ± 1.23 vs. 1.05 ± 1.34 mmHg), daytime systolic blood pressure (-3.61 ± 1.26 vs. 0.80 ± 1.57 mmHg), and nighttime systolic blood pressure $(-3.94 \pm 1.70 \text{ vs. } 1.90 \pm 1.66 \text{ mmHg})$ in prehypertensive obese women, although there were no significant differences in diastolic pressure and other parameters (51).

3.3 Diabetes

Type 1 diabetes, type 2 diabetes, and prediabetes are considered independent risk factors for CVDs. Patients with these conditions

have a 2-4 times higher likelihood of developing CVDs compared to healthy individuals. When CVDs coexist, the risk of major cardiovascular events and all-cause mortality increases (52-55). Improving glycemic and lipid metabolism, controlling blood pressure, and reducing insulin resistance are critical strategies for mitigating the risk of diabetes as a CVD risk factor. A cohort study indicated that overweight/obese type 2 diabetes patients who consume more than 5 grams of green tea daily and have been drinking green tea for over 40 years are associated with a 50% reduction in cardiovascular disease risk (56). Mi et al. (141) discovered that the beneficial effects of EGCG on obesity and diabetes are attributed to its phosphorylation of AMPK and ACC, key enzymes that inhibit lipogenesis. Additionally, EGCG improves disruptions in redox balance and mitochondrial function, alleviating insulin signaling pathway blockages (57). Ren et al. (142), through an intervention in a high-fat diet and streptozotocin-induced type 2 diabetes (T2DM) mouse model, confirmed that EGCG reduced blood glucose levels and improved insulin resistance in T2DM mice. Moreover, EGCG regulated total cholesterol, triglyceride, and low-density lipoprotein receptor levels while reducing lipid deposition in vascular endothelial cells (58). Clearly, tea not only improves blood glucose levels and insulin resistance but also plays a positive role in regulating glucose-lipid metabolism and addressing obesity/ overweight conditions (Figure 4).

Polysaccharides, catechins, and theaflavins in tea are considered primary contributors to its blood glucose-lowering effects, as they improve lipid synthesis, gut microbiota, and glucose transporter proteins (59-61). A randomized controlled trial targeting adults with metabolic syndrome found that the improvement in fasting blood glucose by green tea extract may be attributed to a reduction in blood endotoxins, suppression of intestinal inflammation, and decreased intestinal permeability (62). Yan et al. (143) observed that tea effectively inhibited weight gain, glucose metabolism disorders, and oxidative stress caused by a high-fat diet, while also improving gut microbiota composition (63). Green tea was shown to inhibit NF-κB pathway activation, which upregulated the expression of Glu2 and PPARy, thereby reducing lipid synthesis and promoting glucose transport (64). Theaflavins, at non-cytotoxic doses, significantly enhanced glucose uptake in insulin-resistant cells. This activity was linked to the upregulation of GLUT4 expression and Akt phosphorylation. Moreover, theaflavins mitigated insulin resistance in hepatocytes induced by free fatty acids, indicating their potential role in improving metabolic abnormalities associated with insulin resistance (65).

Furthermore, Hadeel et al. (144) demonstrated that after three weeks of treatment with 100–200 mg/dL doses of green tea extract, wound-related parameters in the skin of T1DM rats significantly improved within 14 days. The intervention regulated the expression of microRNAs (miR-21, miR-23a, miR-146a, miR-29b) and apoptotic genes (Bax, caspase-3, BcL-2). These findings provide potential insights into the role of tea polyphenols in ameliorating apoptosis and promoting angiogenesis (66).

3.4 Obesity

Obesity is widely recognized as a significant risk factor for cardiovascular diseases (CVDs). Studies indicate a strong association between obesity and traditional cardiovascular risk factors such as



hypertension, diabetes, and dyslipidemia, with obese individuals having markedly higher rates of these conditions compared to those of normal weight. Obesity negatively impacts key risk factor levels and mechanisms, increasing the risk of CVDs through mechanisms such as myocardial dysfunction, insulin resistance, inflammation, and abnormal metabolism and hormone secretion by adipose tissue (67–69).

A meta-analysis involving 3,802 participants revealed that green tea extract, as a supplementary intervention for cardiovascular diseases, effectively improved oxidative stress markers such as malondialdehyde and total antioxidant capacity, while also enhancing body composition and obesity-related hormones. Unfortunately, the dose-response evaluation did not yield optimal results, likely due to variations in the quality of evidence included in the analysis (70). Joshua et al. conducted a 12-week intervention among 55 overweight and/or obese adult participants using a multi-component supplement (50 mg forskolin, 500 mg green coffee bean extract, 500 mg green tea extract, 500 mg beetroot extract, 400 mg α -lipoic acid, 200 IU vitamin E, and 200 mg CoQ10) or a placebo. The trial group showed significant reductions in body weight and fat mass, along with improvements in obesity-related plasma markers (GDF15, miR-122, miR-34a), biomarkers for non-alcoholic fatty liver disease (AST, ALT), and resting energy metabolism (71).

Xie et al. (145), in a randomized controlled trial, investigated the effects of a 12-week intervention with decaffeinated green tea polyphenols (400 mg/day) or an equivalent placebo in obese girls aged 6–10. The decaffeinated green tea polyphenol group demonstrated significant reductions in body mass index, body fat percentage, waist circumference, and waist-to-hip ratio, with no adverse effects reported during the study period (72). Tea extracts may exert their effects by

stimulating thermogenesis in brown adipose tissue through the interaction between catechin polyphenols and norepinephrine, which improves body weight and body fat levels (73). Additionally, the efficacy and safety of decaffeinated green tea extract in improving obesity and regulating sex hormone secretion in obese women have been clinically confirmed (74). Related animal studies have shown that the weight control effects of oolong tea extract are dose-dependent and unrelated to dietary changes, with the most significant weight changes observed at moderate doses (75).

3.5 Autonomic function

Cardiac autonomic function is closely related to coronary heart disease, atherosclerosis, and other circulatory system diseases. This may be due to the vascular spasms, platelet aggregation, and activation of inflammatory responses caused by dysregulation of sympathetic and parasympathetic nervous activities, which are common characteristics in the pathology of various heart diseases (76). Studies have confirmed that flavonoid-rich foods such as tea can inhibit excessive sympathetic activity, which may be one of the mechanisms through which these substances exert cardioprotective effects (77).

In vivo studies have demonstrated that black tea significantly increases the high-frequency power and the low-frequency/high-frequency power ratio in the electrocardiograms of rats, improving heart rate variability and enhancing the balance between vagus and sympathetic nerve activity (78). Michelle et al. (146) observed in hypertensive rat models that green tea intervention reduced renal sympathetic nerve activity, improved baroreceptor function, and

significantly lowered blood pressure (79). In vitro research by Fujiko et al. (147) applied various catechin derivatives to the thoracic aortic endothelium of rats. These derivatives inhibited acetylcholine-induced endothelium-dependent relaxation to varying degrees, enhanced vascular reactivity, and exhibited neuromodulatory effects on vascular smooth muscle cells (80).

Heart rate variability (HRV) reflects cardiac autonomic nervous system activity and can quantitatively assess sympathetic and vagal tone, balance, and disease prognosis (81). Reduced HRV is associated with poor cardiovascular outcomes, whereas increased HRV has cardioprotective effects (82). Hinton et al. (148) found that after intervention with γ -aminobutyric acid (GABA)-enriched oolong tea in 30 subjects, autonomic nervous stability and HRV improved, concluding that GABA-enriched oolong tea has beneficial effects on HRV and autonomic nervous function. However, due to the inherent function of GABA and the lack of further mechanistic studies, it remains unclear whether these effects are due to GABA, oolong tea, or their combined action (83).

4 Intervention strategies

The intervention strategies section was developed by setting keywords such as tea, tea extracts, tea components, cardiovascular risk factors, blood glucose, blood lipids, blood pressure, and obesity for searches in PubMed and Web of Science. The document type was limited to clinical studies, systematic reviews, and meta-analyses published between 2010 and 2025. Animal and cell research were excluded from the collected literature. Based on the results of the collected studies, tea intervention strategies were proposed for different cardiovascular risk factors (Figure 5).

In some systematic reviews and meta-analyses, the publication dates of the included studies were relatively early and the intervention dosages maybe unreasonable. To ensure the safety and scientific validity of the recommended strategies, the application dosages also referenced the tea safety standards outlined in the MOFCOMpublished "Technical Guide to Export Commodities."

5 Discussion

This review summarizes the characteristics, chemical components, potential cardiovascular benefits, and possible mechanisms of tea. It also proposes the vision of utilizing tea as a nutritional intervention in primary and secondary prevention of cardiovascular diseases (CVDs). On one hand, tea is a widely consumed soft drink with low cost, offering good economic value, and numerous studies have confirmed the safety of tea interventions in patients with cardiovascular diseases. On the other hand, tea is rich in bioactive compounds such as tea polyphenols, catechins, theanine, and tea polysaccharides. These compounds exhibit multiple effects, including lowering blood lipids, reducing blood pressure, improving diabetes, and providing antioxidant and anti-inflammatory benefits. These attributes align

Cardiovascular Risk Factors	Intervention Strategies			References
	Category	Dosage		References
Lipids	GTE (Cathchins or EGCG)	Cathchins 650-1400mg or EGCG 300- 800mg per day	 ①Some clinical and evidence-based medical studies indicate that black tea has no significant effect on lipids, particularly HDL-C; ②Combining it with aerobic exercise can achieve more optimal results; 	[26][51] [96-103][124,125]
	Black Tea (theaflavins 90mg per Liter)	~600ml per day		
Blood Presure	GTE (EGCG)	EGCG 350-850mg per day	 ①Green tea and GTE are more effective in improving blood pressure compared to black tea; ②When homocysteine levels are elevated, black tea or oolong tea may provide better vascular protection; 	[43][49] [104-113][124,125]
	Green Tea	~1000ml per day		
	Black Tea (Catechin)	~900ml(110-160mg) per day		
Diabetes	GTE (Cathchins or EGCG or Polyphenols)	Cathchins 600-1400mg or EGCG 800mg or 80-240mg per day	 ①Effective on FBG but none for insulin level or HbA1c; ②Lower doses for long-term use of Green Tea and GTE; ③Combining with exercise for better effects; 	[114-125][130]
	BTE (Theaflavins)	Theaflavins 300-1000mg per day		
	Oolong Tea	1500ml with 15g tea leaf per day		
	Green Tea	900ml with 9g tea leaf per day		
Obesity	GTE (EGCG)	450-850mg per day	Combining diet, exercise, and medical interventions can more effectively improve body composition and obesity-related hormone;	[74][126-130]
	Green Tea	800-1500mg per day		

Intervention strategies for cardiovascular risk factors (Green tea extra-GTE, Black tea extra-BTE).

with the needs of primary and secondary prevention of CVDs, making tea a promising choice for nutritional intervention (84–86).

Current tea-related research predominantly focuses on green tea, black tea, and individual component extracts, the former primarily due to their high catechin or theaflavin content, and the latter due to the purity of the extracted components, which facilitates the study of specific bioactivities and mechanisms. Although the application of tea infusions or extracts in studies has increased, better reflecting daily tea consumption, most studies cannot explicitly identify the precise source of tea's therapeutic effects on diseases or symptoms. This may be due to the complex and diverse bioactive components in tea leaves and the still incompletely understood mechanisms of their actions (87).

The chemical composition differences caused by various tea varieties and processing methods add complexity to the research. Therefore, despite the widespread recognition of the health benefits of green and black tea, further in-depth research and exploration are required to clarify tea's therapeutic effects on specific diseases or symptoms. The positive effects of tea components on various diseases are the main focus of most health-related tea research. Chen et al. (149) demonstrated ideal anti-inflammatory effects by inducing the production of nitric oxide, iNOS, COX-2, IL-6, and IL-10 in Raw 264.7 cells using processing techniques of green and black tea applied to coffee leaves. This indicates that the positive effects of tea on human health may not only come from its inherent chemical components but also be influenced by processing methods that alter the content and proportion of active ingredients in tea leaves (88).

The FAO report on tea consumption shows that global tea production and consumption continue to grow, driven by an increasing awareness of tea's health benefits, with black tea and dark tea being the fastest-growing types, primarily in China and other Asian countries (89). Additionally, due to the growth environment and production methods of tea largely originating in Asia and the relatively high number of related studies, research results can vary by region and population (90). For example, the dose-response relationship between green tea consumption and CAD risk shows a significant negative correlation between cardiovascular disease risk and green tea consumption in Asian populations, while this correlation is not observed in Western populations. It was found that an additional cup of tea per day is associated with a 4% reduction in CVD mortality, a 2% reduction in CVD events, and a 1.5% reduction in all-cause mortality, with higher correlations in elderly individuals (91, 92). This situation highlights the lack of broad representativeness in the impact of tea on cardiovascular risk factors. Future research needs to expand the study's geographic scope and include subjects from different racial groups for further exploration.

References

1. Visseren FLJ, Mach F, Smulders YM, Carballo D, Koskinas KC, Bäck M, et al. ESC scientific document group. 2021 ESC guidelines on cardiovascular disease prevention in clinical practice. *Eur J Prev Cardiol.* (2022) 29:5–115. doi: 10.1093/eurjpc/zwab154

2. McPhee PG, Singh S, Morrison KM. Childhood obesity and cardiovascular disease risk: working toward solutions. *Can J Cardiol.* (2020) 36:1352–61. doi: 10.1016/j.cjca.2020.06.020

3. Korczak DJ, Cleverley K, Birken CS, Pignatiello T, Mahmud FH, McCrindle BW. Cardiovascular disease risk factors among children and adolescents with depression. *Front Psych.* (2021) 12:702737. doi: 10.3389/fpsyt.2021.702737

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Funding

The author(s) declare that financial support was received for the research and/or publication of this article. This research is supported by the Hospital Capability Enhancement Project of Xiyuan Hospital of China Academy of Chinese Medical Sciences (Grant No.XYZX0404-03).

Acknowledgments

We would like to express our sincere gratitude to Professor Li Huabin and his team from Sun Yat-sen University for agreeing to the results of their research on the chemical content of the six teas in this paper. Also thanks to Kingsoft Software, the images in this article use materials from their software.

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

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^{4.} Qiu L, Wang W, Sa R, Liu F. Prevalence and risk factors of hypertension, diabetes, and dyslipidemia among adults in Northwest China. *Int J Hypertens*. (2021) 2021:1–10. doi: 10.1155/2021/5528007

^{5.} Paquette M, Bernard S, Paré G, Baass A. Triglycerides, hypertension, and smoking predict cardiovascular disease in dysbetalipoproteinemia. *J Clin Lipidol.* (2020) 14:46–52. doi: 10.1016/j.jacl.2019.12.006

^{6.} Luo Q, Luo L, Zhao J, Wang Y, Luo H. Biological potential and mechanisms of Tea's bioactive compounds: an updated review. *J Adv Res.* (2023) S2090-1232:00378. doi: 10.1016/j.jare.2023.12.004

 Rajendran A, Minhas AS, Kazzi B, Varma B, Choi E, Thakkar A, et al. Sex-specific differences in cardiovascular risk factors and implications for cardiovascular disease prevention in women. *Atherosclerosis.* (2023) 384:117269. doi: 10.1016/j.atherosclerosis.2023.117269

8. Teo KK, Rafiq T. Cardiovascular risk factors and prevention: a perspective from developing countries. *Can J Cardiol*. (2021) 37:733–43. doi: 10.1016/j.cjca.2021.02.009

9. Keller A, Wallace TC. Tea intake and cardiovascular disease: an umbrella review. Ann Med. (2021) 53:929-44. doi: 10.1080/07853890.2021.1933164

10. Tao W, Zhou Z, Zhao B, Wei T. Simultaneous determination of eight catechins and four theaflavins in green, black and oolong tea using new HPLC-MS-MS method. *J Pharm Biomed Anal.* (2016) 131:140–5. doi: 10.1016/j.jpba.2016.08.020

11. Wang W, Liu K, Dong H, Liao W, Yang X, He Q. A frontier exploration of ancient craftsmanship: effects of various tea products in traditional Chinese cuisine "tea flavored beef". *Food Chem.* (2024) 454:139834. doi: 10.1016/j.foodchem.2024.139834

12. Khan N, Mukhtar H. Tea polyphenols in promotion of human health. *Nutrients*. (2018) 11:39. doi: 10.3390/nu11010039

13. Recent insights into the physicochemical properties. Bioactivities and their relationship of tea polysaccharides. *Food Chem.* (2024) 432:137223. doi: 10.1016/j.foodchem.2023.137223

14. Powell-Wiley TM, Poirier P, Burke LE, Després JP, Gordon-Larsen P, Lavie CJ, et al. American Heart Association Council on lifestyle and Cardiometabolic health; council on cardiovascular and stroke nursing; council on clinical cardiology; council on epidemiology and prevention; and stroke council. obesity and cardiovascular disease: a scientific statement from the American Heart Association. *Circulation*. (2021) 143:e984– e1010. doi: 10.1161/CIR.00000000000973

15. Tran HH, Mansoor M, Butt SRR, Satnarine T, Ratna P, Sarker A, et al. Impact of green tea consumption on the prevalence of cardiovascular outcomes: a systematic review. *Cureus*. (2023) 15:e49775. doi: 10.7759/cureus.49775

16. Fang Y, Wang J, Cao Y, Liu W, Duan L, Hu J, et al. The Antiobesity effects and potential mechanisms of Theaflavins. *J Med Food*. (2024) 27:1–11. doi: 10.1089/jmf.2023.K.0180

17. Han LK, Takaku T, Li J, Kimura Y, Okuda H. Anti-obesity action of oolong tea. Int J Obes Relat Metab Disord. (1999) 23:98–105.

18. Zhang S, Takano J, Murayama N, Tominaga M, Abe T, Park I, et al. Subacute ingestion of caffeine and oolong tea increases fat oxidation without affecting energy expenditure and sleep architecture: a randomized, placebo-controlled, double-blinded cross-over trial. *Nutrients*. (2020) 12:3671. doi: 10.3390/nu 12123671

19. Alipour B, Rashidkhani B, Edalati S. Dietary flavonoid intake, total antioxidant capacity and lipid oxidative damage: a cross-sectional study of Iranian women. *Nutrition*. (2016) 32:566–72. doi: 10.1016/j.nut.2015.11.011

20. Raederstorff DG, Schlachter MF, Elste V, Weber P. Effect of EGCG on lipid absorption and plasma lipid levels in rats. *J Nutr Biochem*. (2003) 14:326–32. doi: 10.1016/S0955-2863(03)00054-8

21. Huang F, Zheng X, Ma X, Jiang R, Zhou W, Zhou S, et al. Theabrownin from Puerh tea attenuates hypercholesterolemia via modulation of gut microbiota and bile acid metabolism. *Nat Commun.* (2019) 10:4971. doi: 10.1038/s41467-019-12896-x

22. Kashif M, Imran A, Saeed F, Chatha SAS, Arshad MU. Catechins, theaflavins and ginger freeze-dried extract based functional drink significantly mitigate the hepatic, diabetic and lipid abnormalities in rat model. *Cell Mol Biol (Noisy-le-Grand)*. (2021) 67:132–41. doi: 10.14715/cmb/2021.67.1.20

23. Fu H, He J, Li C, Chang H. Theaflavin-3,3'-Digallate protects liver and kidney functions in diabetic rats by up-regulating Circ-ITCH and Nrf2 signaling pathway. J Agric Food Chem. (2024) 72:14630–9. doi: 10.1021/acs.jafc.3c08251

24. Wang Y, Xia H, Yu J, Sui J, Pan D, Wang S, et al. Effects of green tea catechin on the blood pressure and lipids in overweight and obese population-a meta-analysis. *Heliyon*. (2023) 9:e21228. doi: 10.1016/j.heliyon.2023.e21228

 Hematinezhad Touli M, Elmieh A, Hosseinpour A. The effect of six-week aerobic exercise combined with ease max Incr2Green tea consumption on PON1 and VOand Apelin, blood pressure, and blood lipids reduction in young obese men. Arch Razi Inst. (2022) 77:2115–23. doi: 10.22092/ARI.2022.357847.2109

26. Trautwein EA, Du Y, Meynen E, Yan X, Wen Y, Wang H, et al. Purified black tea theaflavins and theaflavins/catechin supplements did not affect serum lipids in healthy individuals with mildly to moderately elevated cholesterol concentrations. *Eur J Nutr.* (2010) 49:27–35. doi: 10.1007/s00394-009-0045-7

27. Quezada-Fernández P, Trujillo-Quiros J, Pascoe-González S, Trujillo-Rangel WA, Cardona-Müller D, Ramos-Becerra CG, et al. Effect of green tea extract on arterial stiffness, lipid profile and sRAGE in patients with type 2 diabetes mellitus: a randomised, double-blind, placebo-controlled trial. *Int J Food Sci Nutr.* (2019) 70:977–85. doi: 10.1080/09637486.2019.1589430

28. Scolaro B, Nogueira MS, Paiva A, Bertolami A, Barroso LP, Vaisar T, et al. Statin dose reduction with complementary diet therapy: a pilot study of personalized medicine. *Mol Metab.* (2018) 11:137–44. doi: 10.1016/j.molmet.2018.02.005

29. Katada S, Oishi S, Yanagawa K, Ishii S, Oki M, Matsui Y, et al. Concomitant use of tea catechins affects absorption and serum triglyceride-lowering effects of monoglucosyl hesperidin. *Food Funct*. (2021) 12:9339–46. doi: 10.1039/D1FO01917A

30. Doyle AE. Hypertension and vascular disease. *Am J Hypertens*. (1991) 4:103S–6S. doi: 10.1093/ajh/4.2.103S

31. Hodgson JM, Devine A, Puddey IB, Chan SY, Beilin LJ, Prince RL. Tea intake is inversely related to blood pressure in older women. *J Nutr.* (2003) 133:2883–6. doi: 10.1093/jn/133.9.2883

32. Yang YC, Lu FH, Wu JS, Wu CH, Chang CJ. The protective effect of habitual tea consumption on hypertension. *Arch Intern Med.* (2004) 164:1534–40. doi: 10.1001/archinte.164.14.1534

33. Dludla PV, Nkambule BB, Mazibuko-Mbeje SE, Nyambuya TM, Orlando P, Silvestri S, et al. Tea consumption and its effects on primary and secondary prevention of coronary artery disease: qualitative synthesis of evidence from randomized controlled trials. *Clin Nutr ESPEN*. (2021) 41:77–87. doi: 10.1016/j.clnesp.2020.11.006

34. Miller RJ, Jackson KG, Dadd T, Mayes AE, Brown AL, Minihane AM. The impact of the catechol-O-methyltransferase genotype on the acute responsiveness of vascular reactivity to a green tea extract. *Br J Nutr.* (2011) 105:1138–44. doi: 10.1017/S0007114510004836

35. Hodgson JM, Burke V, Puddey IB. Acute effects of tea on fasting and postprandial vascular function and blood pressure in humans. *J Hypertens*. (2005) 23:47–54. doi: 10.1097/00004872-200501000-00012

36. Tinelli C, Di Pino A, Ficulle E, Marcelli S, Feligioni M. Hyperhomocysteinemia as a risk factor and potential nutraceutical target for certain pathologies. *Front Nutr.* (2019) 6:49. doi: 10.3389/fnut.2019.00049

37. Persson M, Josefsson KP, Andersson RG. Tea flavanols inhibit angiotensin converting enzyme activity and increase nitric oxide production in human endothelial cells. *JPharm Pharmacol.* (2006) 58:1139–44. doi: 10.1211/jpp.58.8.0016

38. Rufian-Henares JA, Morales FJ. Angiotensin-I converting enzyme inhibitory activity of coffee melanoidins. *J Agric Food Chem.* (2007) 55:1480–5. doi: 10.1021/jf062604d

39. Hodgson JM. Effects of tea and tea flavonoids on endothelial function and blood pressure: a brief review. *Clin Exp Pharmacol Physiol.* (2006) 33:838–41. doi: 10.1111/j.1440-1681.2006.04450.x

40. Wang L, Tian X. Epigallocatechin-3-Gallate protects against homocysteineinduced brain damage in rats. *Planta Med.* (2018) 84:34–41. doi: 10.1055/s-0043-114865

41. Pei J, Guo S, Zhang C, Xie K. Influence of tea polyphenols on the damage of fibrinolytic system induced by homocysteine in human umbilical vein endothelial cells. *Wei Sheng Yan Jiu.* (2009) 38:47–50. Chinese. doi: 10.19813/j.cnki. weishengyanjiu.2009.01.015

42. Lee WJ, Shim JY, Zhu BT. Mechanisms for the inhibition of DNA methyltransferases by tea catechins and bioflavonoids. *Mol Pharmacol.* (2005) 68:1018–30. doi: 10.1124/mol.104.008367

43. Zhu J, Wang W, Xiong Y, Cooper RS, Du Raza-Arvizu R, Cao G, et al. The association between tea consumption and Hyperhomocysteine in Chinese hypertensive patients. *Am J Hypertens*. (2019) 32:209–15. doi: 10.1093/ajh/hpy163

44. Wang W, Sun Y, Liu J, Wang J, Li Y, Li H, et al. Protective effect of theaflavins on homocysteine-induced injury in HUVEC cells in vitro. *J Cardiovasc Pharmacol.* (2012) 59:434–40. doi: 10.1097/FJC.0b013e318248aeb3

45. Hodgson JM, Burke V, Beilin LJ, Croft KD, Puddey IB. Can black tea influence plasma total homocysteine concentrations? *Am J Clin Nutr.* (2003) 77:907–11. doi: 10.1093/ajcn/77.4.907

46. Dos Passos RR, Santos CV, Priviero F, Briones AM, Tostes RC, Webb RC, et al. Immunomodulatory activity of cytokines in hypertension: a vascular perspective. *Hypertension*. (2024) 81:1411–23. doi: 10.1161/HYPERTENSIONAHA.124.21712

47. Liang Y, Ip MSM, Mak JCW. (–)-Epigallocatechin-3-gallate suppresses cigarette smoke-induced inflammation in human cardiomyocytes via ROS-mediated MAPK and NF- κ B pathways. *Phytomedicine*. (2019) 58:152768. doi: 10.1016/j.phymed.2018.11.028

48. Reddy AT, Lakshmi SP, Maruthi Prasad E, Varadacharyulu NC, Kodidhela LD. Epigallocatechin gallate suppresses inflammation in human coronary artery endothelial cells by inhibiting NF-κB. *Life Sci.* (2020) 258:118136. doi: 10.1016/j.lfs.2020.118136

49. Mahdavi-Roshan M, Salari A, Ghorbani Z, Ashouri A. The effects of regular consumption of green or black tea beverage on blood pressure in those with elevated blood pressure or hypertension: a systematic review and meta-analysis. *Complement Ther Med.* (2020) 51:102430. doi: 10.1016/j.ctim.2020.102430

50. Biesinger S, Michaels HA, Quadros AS, Qian Y, Rabovsky AB, Badger RS, et al. A combination of isolated phytochemicals and botanical extracts lowers diastolic blood pressure in a randomized controlled trial of hypertensive subjects. *Eur J Clin Nutr.* (2016) 70:10–6. doi: 10.1038/ejcn.2015.88

51. Nogueira LP, Nogueira Neto JF, Klein MR, Sanjuliani AF. Short-term effects of green tea on blood pressure, endothelial function, and metabolic profile in obese Prehypertensive women: a crossover randomized clinical trial. *J Am Coll Nutr.* (2017) 36:108–15. doi: 10.1080/07315724.2016.1194236

52. Sattar N, Rawshani A, Franzén S, Rawshani A, Svensson AM, Rosengren A, et al. Age at diagnosis of type 2 diabetes mellitus and associations with cardiovascular and mortality risks. *Circulation*. (2019) 139:2228–37. doi: 10.1161/circulationaha.118.037885

53. Rawshani A, Sattar N, Franzen S, Rawshani A, Hattersley AT, Svensson AM, et al. Excess mortality and cardiovascular disease in young adults with type 1 diabetes in

relation to age at onset: a nationwide, register-based cohort study. Lancet. (2018) 392:477486:477-86. doi: 10.1016/S0140-6736(18)31506-X

54. Rawshani A, Rawshani A, Sattar N, Franzen S, McGuire DK, Eliasson B, et al. Relative prognostic importance and optimal levels of risk factors for mortality and cardiovascular outcomes in type 1 diabetes mellitus. *Circulation*. (2019) 139:19001912. doi: 10.1161/CIRCULATIONAHA.118.037454

55. Livingstone SJ, Looker HC, Hothersall EJ, Wild SH, Lindsay RS, Chalmers J, et al. Risk of cardiovascular disease and total mortality in adults with type 1 diabetes: Scottish registry linkage study. *PLoSMed.* (2012) 9:e1001321. doi: 10.1371/journal.pmed.1001321

56. Liu B, Gu S, Zhang J, Zhou H, Su J, Wang S, et al. Green tea consumption and incidence of cardiovascular disease in type 2 diabetic patients with overweight/obesity: a community-based cohort study. *Arch Public Health.* (2024) 82:18. doi: 10.1186/s13690-024-01242-3

57. Mi Y, Liu X, Tian H, Liu H, Li J, Qi G, et al. EGCG stimulates the recruitment of brite adipocytes, suppresses adipogenesis and counteracts TNF-α-triggered insulin resistance in adipocytes. *Food Funct*. (2018) 9:3374–86. doi: 10.1039/C8FO00167G

58. Ren Z, Yang Z, Lu Y, Zhang R, Yang H. Anti-glycolipid disorder effect of epigallocatechin-3-gallate on high-fat diet and STZ-induced T2DM in mice. *Mol Med Rep.* (2020) 21:2475–83. doi: 10.3892/mmr.2020.11041

59. Esposito F, Pala N, Carcelli M, Boateng ST, D'Aquila PS, Mariani A, et al. α -Glucosidase inhibition by green, white and oolong teas: in vitro activity and computational studies. *J Enzyme Inhib Med Chem.* (2023) 38:2236802. doi: 10.1080/14756366.2023.2236802

60. Liu J, Ding H, Yan C, He Z, Zhu H, Ma KY. Effect of tea catechins on gut microbiota in high fat diet-induced obese mice. *J Sci Food Agric*. (2023) 103:2436–45. doi: 10.1002/jsfa.12476

61. Ni D, Ai Z, Munoz-Sandoval D, Suresh R, Ellis PR, Yuqiong C, et al. Inhibition of the facilitative sugar transporters (GLUTs) by tea extracts and catechins. *FASEB J*. (2020) 34:9995–10010. doi: 10.1096/fj.202000057RR

62. Zeng M, Hodges JK, Pokala A, Khalafi M, Sasaki GY, Pierson J, et al. A green tea extract confection decreases circulating endotoxin and fasting glucose by improving gut barrier function but without affecting systemic inflammation: a double-blind, placebocontrolled randomized trial in healthy adults and adults with metabolic syndrome. *Nutr Res.* (2024) 124:94–110. doi: 10.1016/j.nutres.2024.02.001

63. Liang Y, Wu F, Wu D, Zhu X, Gao X, Hu X, et al. Fu loose tea administration ameliorates obesity in high-fat diet-fed C57BL/6J mice: a comparison with Fu brick tea and orlistat. *Food Secur.* (2024) 13:206. doi: 10.3390/foods13020206

64. Wang R, Gu M, Zhang Y, Zhong Q, Chen L, Li D, et al. Long-term drinking of green tea combined with exercise improves hepatic steatosis and obesity in male mice induced by high-fat diet. *Food Sci Nutr.* (2023) 12:776–85. doi: 10.1002/fsn3.3773

65. Tong T, Ren N, Soomi P, Wu J, Guo N, Kang H, et al. Theaflavins improve insulin sensitivity through regulating mitochondrial biosynthesis in palmitic acid-induced HepG2 cells. *Molecules*. (2018) 23:3382.

66. Al-Rawaf HA, Gabr SA, Alghadir AH. Potential roles of circulating microRNAs in the healing of type 1 diabetic wounds treated with green tea extract: molecular and biochemical study. *Heliyon*. (2023) 9:e22020. doi: 10.1016/j.heliyon.2023.e22020

67. Seifalian AM, Filippatos TD, Joshi J, Mikhailidis DP. Obesity and arterial compliance alterations. *Curr Vasc Pharmacol.* (2010) 8:155–68. doi: 10.2174/157016110790886956

68. Zarzour A, Kim HW, Weintraub NL. Understanding obesity-related cardiovascular disease: It's all about balance. *Circulation*. (2018) 138:64–6. doi: 10.1161/circulationaha.118.034454

69. Rosengren A, Hawken S, Ounpuu S, Sliwa K, Zubaid M, Almahmeed WA, et al. Association of psychosocial risk factors with risk of acute myocardial infarction in 11119 cases and 13648 controls from 52 countries (the INTERHEART study): case-control study. *Lancet.* (2004) 364:953962:953–62. doi: 10.1016/S0140-6736(04)17019-0

70. Vaccarino V, Badimon L, Bremner JD, Cenko E, Cubedo J, Dorobantu M, et al. Position paper of the ESC working group on coronary pathophysiology and microcirculation. *Eur Heart J.* (2018) 2020:16871696. doi: 10.1093/eurheartj /ehy913

71. Nederveen JP, Mastrolonardo AJ, Xhuti D, Di Carlo A, Manta K, Fuda MR, et al. Novel multi-ingredient supplement facilitates weight loss and improves body composition in overweight and obese individuals: a randomized, double-blind, placebocontrolled clinical trial. *Nutrients*. (2023) 15:3693. doi: 10.3390/nu15173693

72. Xie L, Tang Q, Yao D, Gu Q, Zheng H, Wang X, et al. Effect of decaffeinated green tea polyphenols on body fat and precocious puberty in obese girls: a randomized controlled trial. *Front Endocrinol.* (2021) 12:736724.

73. Asbaghi O, Rezaei Kelishadi M, Larky DA, Bagheri R, Amirani N, Goudarzi K, et al. The effects of green tea extract supplementation on body composition, obesityrelated hormones and oxidative stress markers: a grade-assessed systematic review and dose-response meta-analysis of randomised controlled trials. *Br J Nutr.* (2024) 131:1125–57. doi: 10.1017/S000711452300260X

74. Dulloo AG, Seydoux J, Girardier L, Chantre P, Vandermander J. Green tea and thermogenesis: interactions between catechin-polyphenols, caffeine and sympathetic activity. *Int J Obes Relat Metab Disord.* (2000) 24:252–8. doi: 10.1038/sj.ijo. 0801101

75. Tung YC, Liang ZR, Yang MJ, Ho CT, Pan MH. Oolong tea extract alleviates weight gain in high-fat diet-induced obese rats by regulating lipid metabolism and modulating gut microbiota. *Food Funct.* (2022) 13:2846–56. doi: 10.1039/D1FO03356E

76. Xie L, Tang Q, Yao D, Gu Q, Zheng H, Wang X, et al. Effect of decaffeinated green tea polyphenols on body fat and Precocio us puberty in obese girls: a randomized controlled trial. *Front Endocrinol.* (2021) 12:736724. doi: 10.3389/fendo.2021.736724

77. Braun J, Patel M, Kameneva T, Keatch C, Lambert G, Lambert E. Central stress pathways in the development of cardiovascular disease. *Clin Auton Res.* (2023) 34:99–116. doi: 10.1007/s10286-023-01008-x

78. Bruno RM, Ghiadoni L. Polyphenols, antioxidants and the sympathetic nervous system. *Curr Pharm Des.* (2018) 24:130–9. doi: 10.2174/1381612823666171114170642

79. Garcia ML, Pontes RB, Nishi EE, Ibuki FK, Oliveira V, Sawaya AC, et al. The antioxidant effects of green tea reduces blood pressure and sympathoexcitation in an experimental model of hypertension. *J Hypertens*. (2017) 35:348–54. doi: 10.1097/HJH.00000000001149

80. Sanae F, Miyaichi Y, Kizu H, Hayashi H. Effects of catechins on vascular tone in rat thoracic aorta with endothelium. *Life Sci.* (2002) 71:2553–62. doi: 10.1016/S0024-3205(02)02080-5

81. Cygankiewicz I, Zareba W. Heart rate variability. *Handb Clin Neurol.* (2013) 117:379–93. doi: 10.1016/B978-0-444-53491-0.00031-6

82. Rajendra Acharya U, Paul Joseph K, Kannathal N, Lim CM, Suri JS. Heart rate variability: a review. *Med Biol Eng Comput.* (2006) 44:1031–51. doi: 10.1007/s11517-006-0119-0

83. Hinton T, Jelinek HF, Viengkhou V, Johnston GA, Matthews S. Effect of GABAfortified oolong tea on reducing stress in a university student cohort. *Front Nutr.* (2019) 6:27. doi: 10.3389/fnut.2019.00027

84. Sagris M, Vlachakis PK, Simantiris S, Theofilis P, Gerogianni M, Karakasis P, et al. From a cup of tea to cardiovascular care: vascular mechanisms of action. *Life*. (2024) 14:1168. doi: 10.3390/life14091168

85. American Association of Cardiovascular & Pulmonary Rehabilitation. Guidelines for cardiac rehabilitation and secondary prevention programs. US: Promoting Health & Preventing Disease Human Kinetics Publishers (1999).

86. Thomas RJ, Huang HH. Cardiac rehabilitation for secondary prevention of cardiovascular disease: 2019 update. *Curr Treat Options Cardiovasc Med.* (2019) 21:56. doi: 10.1007/s11936-019-0759-7

87. Ho C-T, Lin J-K, Shahidi F. Tea and tea products: Chemistry and health-promoting properties. *1st* ed. Boca Raton: CRC Press (2008).

88. Chen X, Mu K, Kitts DD. Characterization of phytochemical mixtures with inflammatory modulation potential from coffee leaves processed by green and black tea processing methods. *Food Chem.* (2019) 271:248–58. doi: 10.1016/j.foodchem. 2018.07.097

89. United Nations FAO. International tea market: Market situation, prospects and emerging issues. (2022). Available at: https://openknowledge.fao.org/server/api/core/bitstreams/e1d8588a-ddba-4b49-9897-311611391a76/content

90. Peters U, Poole C, Arab L. Does tea affect cardiovascular disease? A meta-analysis. *Am J Epidemiol.* (2001) 154:495–503. doi: 10.1093/aje/154.6.495

91. Yang X, Dai H, Deng R, Zhang Z, Quan Y, Giri M, et al. Association between tea consumption and prevention of coronary artery disease: a systematic review and dose-response meta-analysis. *Front Nutr.* (2022) 9:1021405. doi: 10.3389/fnut.2022.1021405

92. Chung M, Zhao N, Wang D, Shams-White M, Karlsen M, Cassidy A, et al. Doseresponse relation between tea consumption and risk of cardiovascular disease and allcause mortality: a systematic review and Meta-analysis of population-based studies. *Adv Nutr.* (2020) 11:790–814. doi: 10.1093/advances/nmaa010

93. Tang GY, Zhao CN, Xu XY, Gan RY, Cao SY, Liu Q, et al. Phytochemical composition and antioxidant capacity of 30 Chinese teas. *Antioxidants*. (2019) 8:180. doi: 10.3390/antiox8060180

94. Mirza O, Henriksen A, Ostergaard L, Welinder KG, Gajhede M. Arabidopsis thaliana peroxidase N: structure of a novel neutral peroxidase. Acta Crystallogr D Biol Crystallogr. (2000) 56:372–5. doi: 10.1107/S0907444999016340

95. Putnam CD, Arvai AS, Bourne Y, Tainer JA. Active and inhibited human catalase structures: ligand and NADPH binding and catalytic mechanism. *J Mol Biol.* (2000) 296:295–309.

96. Yuan F, Dong H, Fang K, Gong J, Lu F. Effects of green tea on lipid metabolism in overweight or obese people: a meta-analysis of randomized controlled trials. *Mol Nutr Food Res.* (2018) 62. doi: 10.1002/mnfr.201601122

97. Xu R, Yang K, Li S, Dai M, Chen G. Effect of green tea consumption on blood lipids: a systematic review and meta-analysis of randomized controlled trials. *Nutr J.* (2020) 19:48. doi: 10.1186/s12937-020-00557-5

98. Li A, Wang Q, Li P, Zhao N, Liang Z. Effects of green tea on lipid profile in overweight and obese women. *Int J Vitam Nutr Res.* (2024) 94:239–51. doi: 10.1024/0300-9831/a000783

99. Kim A, Chiu A, Barone MK, Avino D, Wang F, Coleman CI, et al. Green tea catechins decrease total and low-density lipoprotein cholesterol: a systematic review and meta-analysis. *J Am Diet Assoc.* (2011) 111:1720–9. doi: 10.1016/j.jada.2011.08.009

100. Bahorun T, Luximon-Ramma A, Neergheen-Bhujun VS, Gunness TK, Googoolye K, Auger C, et al. The effect of black tea on risk factors of cardiovascular disease in a normal population. *Prev Med.* (2012) 54:S98–S102. doi: 10.1016/j.ypmed.2011.12.009

101. Macêdo APA, Gonçalves MDS, Barreto Medeiros JM, David JM, Villarreal CF, Macambira SG, et al. Potential therapeutic effects of green tea on obese lipid profile - a systematic review. *Nutr Health.* (2022) 28:401–15. doi: 10.1177/026010602110 73236

102. Araya-Quintanilla F, Gutiérrez-Espinoza H, Moyano-Gálvez V, Muñoz-Yánez MJ, Pavez L, García K. Effectiveness of black tea versus placebo in subjects with hypercholesterolemia: a PRISMA systematic review and meta-analysis. *Diabetes Metab Syndr.* (2019) 13:2250–8. doi: 10.1016/j.dsx.2019.05.019

103. Wang D, Chen C, Wang Y, Liu J, Lin R. Effect of black tea consumption on blood cholesterol: a meta-analysis of 15 randomized controlled trials. *PLoS One.* (2014) 9:e107711. doi: 10.1371/journal.pone.0107711

104. Liu G, Mi XN, Zheng XX, Xu YL, Lu J, Huang XH. Effects of tea intake on blood pressure: a meta-analysis of randomised controlled trials. *Br J Nutr*. (2014) 112:1043–54. doi: 10.1017/S0007114514001731

105. Li G, Zhang Y, Thabane L, Mbuagbaw L, Liu A, Levine MA, et al. Effect of green tea supplementation on blood pressure among overweight and obese adults: a systematic review and meta-analysis. *J Hypertens.* (2015) 33:243–54. doi: 10.1097/HJH.00000000000426

106. Yarmolinsky J, Gon G, Edwards P. Effect of tea on blood pressure for secondary prevention of cardiovascular disease: a systematic review and meta-analysis of randomized controlled trials. *Nutr Rev.* (2015) 73:236–46. doi: 10.1093/nutrit/ nuv001

107. Xu R, Yang K, Ding J, Chen G. Effect of green tea supplementation on blood pressure: a systematic review and meta-analysis of randomized controlled trials. *Medicine*. (2020) 99:e19047. doi: 10.1097/MD.000000000019047

108. Onakpoya I, Spencer E, Heneghan C, Thompson M. The effect of green tea on blood pressure and lipid profile: a systematic review and meta-analysis of randomized clinical trials. *Nutr Metab Cardiovasc Dis.* (2014) 24:823–36. doi: 10.1016/j.numecd.2014.01.016

109. Grassi D, Draijer R, Desideri G, Mulder T, Ferri C. Black tea lowers blood pressure and wave reflections in fasted and postprandial conditions in hypertensive patients: a randomised study. *Nutrients*. (2015) 7:1037–51. doi: 10.3390/nu70 21037

110. Al-Shafei AIM, El-Gendy OAA. Regular consumption of green tea improves pulse pressure and induces regression of left ventricular hypertrophy in hypertensive patients. *Physiol Rep.* (2019) 7:e14030. doi: 10.14814/phy2.14030

111. Quan J, Zhang T, Gu Y, Meng G, Zhang Q, Liu L, et al. Green tea intake and the risk of hypertension in premenopausal women: the TCLSIH cohort study. *Food Funct.* (2023) 14:4406–13. doi: 10.1039/D2FO03342A

112. Khalesi S, Sun J, Buys N, Jamshidi A, Nikbakht-Nasrabadi E, Khosravi-Boroujeni H. Green tea catechins and blood pressure: a systematic review and meta-analysis of randomised controlled trials. *Eur J Nutr.* (2014) 53:1299–311. doi: 10.1007/s00394-014-0720-1

113. Lin CC, Hsieh CY, Chen LF, Chen YC, Ho TH, Chang SC, et al. Versatile effects of GABA oolong tea on improvements in diastolic blood pressure, alpha brain waves, and quality of life. *Food Secur*. (2023) 12:4101. doi: 10.3390/foods12224101

114. Jia MJ, Liu XN, Liang YC, Liu DL, Li HL. The effect of green tea on patients with type 2 diabetes mellitus: a meta-analysis. *Medicine (Baltimore)*. (2024) 103:e39702. doi: 10.1097/MD.000000000039702

115. Li X, Wang W, Hou L, Wu H, Wu Y, Xu R, et al. Does tea extract supplementation benefit metabolic syndrome and obesity? A systematic review and meta-analysis. *Clin Nutr.* (2020) 39:1049–58. doi: 10.1016/j.clnu.2019.05.019

116. Li Y, Wang C, Huai Q, Guo F, Liu L, Feng R, et al. Effects of tea or tea extract on metabolic profiles in patients with type 2 diabetes mellitus: a meta-analysis of ten randomized controlled trials. *Diabetes Metab Res Rev.* (2016) 32:2–10. doi: 10.1002/dmrr.2641

117. Xu R, Bai Y, Yang K, Chen G. Effects of green tea consumption on glycemic control: a systematic review and meta-analysis of randomized controlled trials. *Nutr Metab.* (2020) 17:56. doi: 10.1186/s12986-020-00469-5

118. Yang CY, Yen YY, Hung KC, Hsu SW, Lan SJ, Lin HC. Inhibitory effects of pu-erh tea on alpha glucosidase and alpha amylase: a systemic review. *Nutr Diabetes.* (2019) 9:23. doi: 10.1038/s41387-019-0092-y

119. Momose Y, Maeda-Yamamoto M, Nabetani H. Systematic review of green tea epigallocatechin gallate in reducing low-density lipoprotein cholesterol levels of humans. *Int J Food Sci Nutr.* (2016) 67:606–13. doi: 10.1080/09637486.2016.1196655

120. Asbaghi O, Fouladvand F, Gonzalez MJ, Ashtary-Larky D, Choghakhori R, Abbasnezhad A. Effect of green tea on glycemic control in patients with type 2 diabetes mellitus: a systematic review and meta-analysis. *Diabetes Metab Syndr*. (2021) 15:23–31. doi: 10.1016/j.dsx.2020.11.004

121. Wang X, Tian J, Jiang J, Li L, Ying X, Tian H, et al. Effects of green tea or green tea extract on insulin sensitivity and glycaemic control in populations at risk of type 2 diabetes mellitus: a systematic review and meta-analysis of randomised controlled trials. *J Hum Nutr Diet.* (2014) 27:501–12. doi: 10.1111/jhn.12181

122. He M, Lyu X. Application of BRAFO-tiered approach for health benefit-risk assessment of dark tea consumption in China. *Food Chem Toxicol.* (2021) 158:112615. doi: 10.1016/j.fct.2021.112615

123. Chen S, Peng D, Shan Y, Liu F, Du R, Bao Y, et al. Black tea drinks with inulin and dextrin reduced postprandial plasma glucose fluctuations in patients with type 2 diabetes: an acute, randomized, placebo-controlled, single-blind crossover study. *Nutr Diabetes*. (2024) 14:95. doi: 10.1038/s41387-024-00351-w

124. Zamani M, Kelishadi MR, Ashtary-Larky D, Amirani N, Goudarzi K, Torki IA, et al. The effects of green tea supplementation on cardiovascular risk factors: a systematic review and meta-analysis. *Front Nutr.* (2023) 9:1084455. doi: 10.3389/fnut.2022.1084455

125. Neyestani TR, Nikooyeh B. A comprehensive overview on the effects of green tea on anthropometric measures, blood pressure, glycemic and lipidemic status: an umbrella review and meta meta-analysis study. *Nutr Metab Cardiovasc Dis.* (2022) 32:2026–40. doi: 10.1016/j.numecd.2022.05.021

126. Zhang Y, Tang N, Xia W, Sanjid Seraj S, Pereira M, Velu P, et al. The effect of green tea supplementation on the anthropometric outcomes in overweight and obese women: a time and dose-response meta-analysis of randomized controlled trials. *Crit Rev Food Sci Nutr.* (2024) 64:10138–47. doi: 10.1080/10408398.2023.2220796

127. Lin Y, Shi D, Su B, Wei J, Găman MA, Sedanur Macit M, et al. The effect of green tea supplementation on obesity: a systematic review and dose-response metaanalysis of randomized controlled trials. *Phytother Res.* (2020) 34:2459–70. doi: 10.1002/ptr.6697

128. Baladia E, Basulto J, Manera M, Martínez R, Calbet D. Efecto del consumo de té verde o extractos de té verde en el peso y en la composición corporal; revisión sistemática y metaanálisis [effect of green tea or green tea extract consumption on body weight and body composition; systematic review and meta-analysis]. *Nutr Hosp.* (2014) 29:479–90. doi: 10.3305/nh.2014.29.3.7118

129. Vázquez Cisneros LC, López-Uriarte P, López-Espinoza A, Navarro Meza M, Espinoza-Gallardo AC, Guzmán Aburto MB. Efectos del té verde y su contenido de galato de epigalocatequina (EGCG) sobre el peso corporal y la masa grasa en humanos. Una revisión sistemática [effects of green tea and its epigallocatechin (EGCG) content on body weight and fat mass in humans: a systematic review]. *Nutr Hosp.* (2017) 34:731–7. doi: 10.20960/nh.753

130. Asbaghi O, Fouladvand F, Gonzalez MJ, Aghamohammadi V, Choghakhori R, Abbasnezhad A. Effect of green tea on anthropometric indices and body composition in patients with type 2 diabetes mellitus: a systematic review and Meta-analysis. *Complement Med Res.* (2021) 28:244–51. doi: 10.1159/000511665

131. Fu H, He J, Li C, Chang H. Theaflavin-3,3'-Digallate Protects Liver and Kidney Functions in Diabetic Rats by Up-Regulating Circ-ITCH and Nrf2 Signaling Pathway. J Agric Food Chem. (2024) 72:14630–9.

132. Trautwein EA, Du Y, Meynen E, Yan X, Wen Y, Wang H, et al. Purified black tea theaflavins and theaflavins/catechin supplements did not affect serum lipids in healthy individuals with mildly to moderately elevated cholesterol concentrations. *Eur J Nutr.* (2010) 49:27–35.

133. Quezada-Fernández P, Trujillo-Quiros J, Pascoe-González S, Trujillo-Rangel WA, Cardona-Müller D, Ramos-Becerra CG, et al. Effect of green tea extract on arterial stiffness, lipid profile and sRAGE in patients with type 2 diabetes mellitus: a randomised, double-blind, placebo-controlled trial. *Int J Food Sci Nutr.* (2019) 70:977–85.

134. Scolaro B, Nogueira MS, Paiva A, Bertolami A, Barroso LP, Vaisar T, et al. Statin dose reduction with complementary diet therapy: A pilot study of personalized medicine. *Mol Metab.* (2018) 11:137–44.

135. Katada S, Oishi S, Yanagawa K, Ishii S, Oki M, Matsui Y, et al. Concomitant use of tea catechins affects absorption and serum triglyceride-lowering effects of monoglucosyl hesperidin. *Food Funct.* (2021) 12:9339–46.

136. Hodgson JM, Devine A, Puddey IB, Chan SY, Beilin LJ, Prince RL. Tea intake is inversely related to blood pressure in older women. *J Nutr.* (2003) 133:2883–6.

137. Wang L, Tian X. Epigallocatechin-3-Gallate Protects against Homocysteine-Induced Brain Damage in Rats. *Planta Med.* (2018) 84:34–41.

138. Reddy AT, Lakshmi SP, Maruthi Prasad E, Varadacharyulu NC, Kodidhela LD. Epigallocatechin gallate suppresses inflammation in human coronary artery endothelial cells by inhibiting NF-κB. *Life Sci.* (2020) 258:118136

139. Mahdavi-Roshan M, Salari A, Ghorbani Z, Ashouri A. The effects of regular consumption of green or black tea beverage on blood pressure in those with elevated blood pressure or hypertension: A systematic review and meta-analysis. *Complement Ther Med.* (2020) 51:102430

140. Biesinger S, Michaels HA, Quadros AS, Qian Y, Rabovsky AB, Badger RS, et al. A combination of isolated phytochemicals and botanical extracts lowers diastolic blood pressure in a randomized controlled trial of hypertensive subjects. *Eur J Clin Nutr.* (2016) 70:10–6.

141. Mi Y, Liu X, Tian H, Liu H, Li J, Qi G, et al. EGCG stimulates the recruitment of brite adipocytes, suppresses adipogenesis and counteracts TNF- α -triggered insulin resistance in adipocytes. *Food Funct*. (2018) 9:3374–86.

142. Ren Z, Yang Z, Lu Y, Zhang R, Yang H. Anti-glycolipid disorder effect of epigallocatechin-3-gallate on high-fat diet and STZ-induced T2DM in mice. *Mol Med Rep.* (2020) 21:2475–83.

143. Liang Y, Wu F, Wu D, Zhu X, Gao X, Hu X, et al. Fu Loose Tea Administration Ameliorates Obesity in High-Fat Diet-Fed C57BL/6J Mice: A Comparison with Fu Brick Tea and Orlistat. *Foods.* (2024) 13:206.

144. Al-Rawaf HA, Gabr SA, Alghadir AH. Potential roles of circulating microRNAs in the healing of type 1 diabetic wounds treated with green tea extract: molecular and biochemical study. *Heliyon*. (2023) 9:e22020

145. Xie L, Tang Q, Yao D, Gu Q, Zheng H, Wang X, et al. Effect of Decaffeinated Green Tea Polyphenols on Body Fat and Precocious Puberty in Obese Girls: A

Randomized Controlled Trial. Front Endocrinol (Lausanne). (2021) 12:736724

146. Garcia ML, Pontes RB, Nishi EE, Ibuki FK, Oliveira V, Sawaya AC, et al. The antioxidant effects of green tea reduces blood pressure and sympathoexcitation in an experimental model of hypertension. *J Hypertens.* (2017) 35:348–54.

147. Sanae F, Miyaichi Y, Kizu H, Hayashi H. Effects of catechins on vascular tone in rat thoracic aorta with endothelium. *Life Sci.* (2002) 71:2553–62.

148. Hinton T, Jelinek HF, Viengkhou V, Johnston GA, Matthews S. Effect of GABA-Fortified Oolong Tea on Reducing Stress in a University Student Cohort. *Front Nutr.* (2019) 6:27.

149. Chen X, Mu K, Kitts DD. Characterization of phytochemical mixtures with inflammatory modulation potential from coffee leaves processed by green and black tea processing methods. *Food Chem.* (2019) 271:248–58.